Application No. Not Yet Assigned Docket No.: 66015(71432)
Amendment dated July 28, 2006

First Preliminary Amendment Page 3

AMENDMENTS TO THE CLAIMS

Claims 1-29 (cancelled)

30. A modified siRNA comprising a sense strand and an antisense strand, wherein the

sense strand comprises a modified RNA nucleotide in at least one of positions 8-14,

calculated from the 5'-end.

31. The modified siRNA according to claim 30, wherein the sense strand comprises a

modified RNA nucleotide in at least one position selected from the group consisting of

position 8, position 9, position 10, position 11, position 12, position 13 and position 14,

calculated from the 5'-end.

32. The modified siRNA according to claim 30, wherein said modified RNA nucleotide

comprises a modification selected from the group consisting of a non-RNA nucleobase,

a sugar moiety which differs from ribose, an internucleoside linkage group which differs

from phosphate, and combinations thereof.

33. The modified siRNA according to claim 32, wherein said modified RNA nucleotide

comprises a non-RNA nucleobase.

34. The modified siRNA according to claim 33, wherein the sense strand comprises a

non-RNA nucleobase in at least one position selected from the group consisting of

position 9, position 10, position 11, position 12 and position 13, calculated from the 5'-

end.

The modified siRNA according to claim 34, wherein the sense strand comprises a

non-RNA nucleobase in a position selected from the group consisting of position 10,

position 11 and both of positions 10 and 11, calculated from the 5'-end.

36. The modified siRNA according to claim 35, wherein the sense strand comprises a

non-RNA nucleobase in position 10, calculated from the 5'-end.

Application No. Not Yet Assigned Amendment dated July 28, 2006 First Preliminary Amendment Page 4

- 37. The modified siRNA according to claim 33, wherein said non-RNA nucleobase is selected from the group consisting of thymine (T), 5-methylcytosine (MeC), isocytosine, pseudoisocytosine, 5-bromouracil, 5-propynyluracil, 5-propyny-6-fluoroluracil, 5-methylthiazoleuracil, 6-aminopurine, 2-aminopurine, inosine, 2,6-diaminopurine, 7-propyne-7-deazaadenine, 7-propyne-7-deazaguanine and 2-chloro-6-aminopurine.
- 38. The modified siRNA according to claim 37, wherein said non-RNA nucleobase is T or ^{Me}C.
- 39. The modified siRNA according to claim 38, wherein said non-RNA nucleobase is T.
- 40. The modified siRNA according to claim 32, wherein said modified RNA nucleotide comprises a sugar moiety which differs from ribose.
- 41. The modified siRNA according to claim 40, wherein the sense strand comprises a sugar moiety which differs from ribose, in at least one position selected from the group consisting of position 9, position 10, position 11, position 12 and position 13, calculated from the 5'-end.
- 42. The modified siRNA according to claim 41, wherein the sense strand comprises a sugar moiety which differs from ribose, in at least one position selected from the group consisting of position 11, position 12 and position 13, calculated from the 5'-end.
- 43. The modified siRNA according to claim 42, wherein the sense strand comprises a sugar moiety which differs from ribose in position 12, calculated from the 5'-end.
- 44. The modified siRNA according to claim 40, wherein said sugar moiety differs from ribose in that the ribose 2'-OH group has been modified.
- 45. The modified siRNA according to claim 44, wherein said sugar moiety is LNA.

Application No. Not Yet Assigned Docket No.: 66015(71432)
Amendment dated July 28, 2006

First Preliminary Amendment Page 5

46. The modified siRNA according to claim 45, wherein said LNA is selected from the

group consisting of thio-LNA, amino-LNA, oxy-LNA and ena-LNA.

47. The modified siRNA according to claim 46, wherein said LNA is oxy-LNA.

48. The modified siRNA according to claim 47, wherein said oxy-LNA is in the beta-D

form.

49. The modified siRNA according to claim 32, wherein said modified RNA nucleotide

comprises an internucleoside linkage group which differs from phosphate.

50. The modified siRNA according to claim 49, wherein the sense strand comprises an

internucleoside linkage group which differs from phosphate, in at least one position

selected from the group consisting of position 9, position 10, position 11, position 12

and position 13, calculated from the 5'-end.

51. The modified siRNA according to claim 50, wherein the sense strand comprises an

internucleoside linkage group which differs from phosphate, in at least one position

selected from the group consisting of position 11, position 12 and position 13,

calculated from the 5'-end.

52. The modified siRNA according to claim 51, wherein the sense strand comprises an

internucleoside linkage group which differs from phosphate in position 12, calculated

from the 5'-end.

53. The modified siRNA according to claim 32, wherein said internucleoside linkage

group which differs from phosphate is phosphorothioate (-O-P(O,S)-O-).

54. The modified siRNA according to claim 30, wherein at least one of the strands

further comprises at least one modified RNA nucleotide.

Docket No.: 66015(71432)

Application No. Not Yet Assigned Amendment dated July 28, 2006 First Preliminary Amendment

Page 6

55. The modified siRNA according to claim 32 wherein the sense strand comprises a

sugar moiety which differs from ribose in position 12, calculated from the 5'-end, and

wherein the sense strand comprises a non-RNA nucleobase in position 10, calculated

from the 5'-end.

56. The modified siRNA according to claim 30, wherein each strand comprises 12-35

monomers.

57. A pharmaceutical composition comprising the modified siRNA as

defined in claim 1 and a pharmaceutically acceptable diluent, carrier or

adjuvant.